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Aug 20, 1986

DERWENT-ACC-NO: 1986-219977

DERWENT-WEEK: 198634

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TITLE: Stabilised active forms of vitamin=D3 - contg. neutral or basic amino acid stabiliser

INVENTOR: IGUSA, K; NEMOTO, K ; OGASAWARA, T

PATENT-ASSIGNEE:

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CODE

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PRIORITY-DATA: 1985JP-0028073 (February 14, 1985)

PATENT-FAMILY:

PUB-NO	PUB-DATE	LANGUAGE	PAGES	MAIN-IPC
EP 191489 A	August 20, 1986	E	010	
DE 3680000 G	August 8, 1991		000	
EP 191489 B	July 3, 1991		000	
JP 62000017 A	January 6, 1987		000	
US 4836957 A	June 6, 1989		000	

DESIGNATED-STATES: DE FR GB DE FR GB

CITED-DOCUMENTS:A3...8734; CH 304182 ; EP 132821 ; EP 144434 ; GB 2052976 ; GB 799068 ; No-SR.Pub ; WO 8404244

APPLICATION-DATA:

PUB-NO	APPL-DATE	APPL-NO	DESCRIPTOR
EP 191489A	February 13, 1986	1986EP-0101828	
JP62000017A	January 31, 1986	1986JP-0018047	
US 4836957A	February 11, 1986	1986US-0828596	

INT-CL (IPC): A61K 31/59; A61K 47/00; C07J 9/00

ABSTRACTED-PUB-NO: EP 191489A

BASIC-ABSTRACT:

Prepns. contg. an active form (I) of vitamin D3 are stabilised by incorporation of an amino acid (II), contg. neither an S atom nor an acid amide gp. and which is neither an acidic amino acid nor a salt of a basic amino acid. Pref. (II) is neutral, e.g. alanine, valine, proline, phenylalanine, tryptophan, leucine, isoleucine, glycine or serine, or basic, e.g. lysine, arginine or histidine. Pref. (I) is 1alpha-hydroxyvitamin D3, 1alpha,24 (or 25)-dihydroxyvitamin D3, 1alpha,24,25-trihydroxyvitamin D3, 26,27 -hexafluoro-1alpha,25-dihydro xyvitamin D3, or 25-hydroxyvitamin D3.

USE/ADVANTAGE - (I) improve Ca metabolism, esp. bone metabolism, and are useful

in treatment of bone diseases resulting from impaired vitamin D activity in the kidney, as well as in treatment of diseases difficult to cure by conventional vitamin D3 drugs, e.g. chronic renal insufficiency, hypoparathyroidism, vitamin D-resistant rickets and osteomalacia. (I) are also admin. to premature babies to sustain their growth. The presence of (II) improves the storage stability of preps. contg.

ABSTRACTED-PUB-NO:

EP 191489B

EQUIVALENT-ABSTRACTS:

A prep. containing an active form of vitamin D3 stabilised by incorporation of an amino acid that is neither one containing a sulphur atom or an acid amide gp. in its structure, nor an acidic amino acid nor a salt of a basic amino acid, except the solid dietary mixt. having the following compsn.: 7.824 g 1-amino acids (alanine, arginine, asparagine, glutamine, glycocoll, histidine hydrochloride, isoleucine, leucine, lysine hydrochloride, methionine, monomagnesium-1-glu tamate, phenylalanine, proline, serine, threonine, tryptophan, tyrosine ethyl ester hydrochloride, valine); 0.33 g neutral oil (MCT-medium chain triglycerides); 2.34 g sunflower oil 2.00 g a proportion of essential fatty acids; 1.2 g carnitine; 74.0 g carbohydrates (maltodextrin e, cold swelling starch, SCChROSQ); 14. m gO ChlOriQ 4. ClCium Chlroi chloride; 726.0 mg dipotassium hydrogen phosphate; 633.0 mg calcium glycerophosphate; 0.387 mg copper acetate (1H2O); 28.8 mg iron-II-gluconate (2H2O) 53.3 micro g potassium iodide; 3.33 mg manganese acetate (4H2O); 2.0 mg zinc oxide vitamins; 16.00 mg ascorbic acid (C); 40.00 micro g biotin; 295.00 mg calcium phosphoryl choline chloride (4H2O); 100.00 micro g folic acid; 3.30 mg nicotinic acid amide; 2.50 mg calcium-d-pantothenate; 330.30 micro g riboflavine-5'- phosphoric acid ester, sodium salt (B2); 333.30 micro g aneurine nitrate (B1); 333.30 micro g pyridoxine hydrochloride (B6); 1.00 g cyanocobalamine (vitamin B12 cyanocomplex); 0.83 mg vitamin A acetate, oily concentrate (1g=10 power 6 IU); 5.00 mg alpha-tocopherol acetate (vitamin E acetate); 3.5 g cholecalciferol cholesterol (D3 cholesterol); 40.0 mg myoinositol. (10pp)

US 4836957A

Prepn. comprises active form of vitamin D3 viz. 1 alpha-hydroxy vitamin D3, 1 alpha 24- and 25-dihydroxy-, 1 alpha, 24 and 25-trihydroxy-, 26, 27-hexafluoro-1 alpha-, 25-dihydroxy-, 24-hydroxy-, and 25-hydroxy-vit amin D3; and stabilizing amt. basic or neutral amino acid free of acid amide gps. and S. Amt. may be 10-500 times wt. vit D3 for liq. preps. and 5000-50000 for solid preps.

Pref. amino acids are Ala, Val, Pro, Phe Trp. Leu, i-Leu, Gly, Ser, Lys, Arg or His, salts and N-phenyl derivs.

USE/ADVANTAGE - Solubilization and stabilization of active forms of vitamin D3 to improve Ca and bone metabolism, esp. during blood dialysis, for premature babies and treatment of difficult forms of vitamin D3 deficiency. (4pp)t

TITLE-TERMS : STABILISED ACTIVE FORM VITAMIN=D3 CONTAIN NEUTRAL BASIC AMINO ACID STABILISED

DERWENT-CLASS: B01 B05 E15

CPI-CODES: B03-G; B12-G03; B12-G06; B12-J01; B12-J08; B12-M06; E03; E06-D01; E07-D03; E07-D09B; E10-A17; E10-B02D3; E10-B02D4; E10-B02D6; E10-E04J;

CHEMICAL-CODES:

Chemical Indexing M2 *01*

Fragmentation Code

H1 H100 H181 J0 J011 J1 J171 M280 M312 M321
M331 M340 M342 M349 M381 M391 M416 M431 M620 M782
M903 M910 Q620